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Practitioner's Docket No. MPI95-015P1RCPA1DV1M

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:	Phillips, David R., et al		
Application No.:	09/801,089	Group No.:	1644
Filed:	March 8, 2001	Examiner:	Ewoldt, Gerald R.
For:	MODULATION OF INTEGRIN-MEDIATED SIGNAL TRANSDUCTION		

Mail Stop DD
Commissioner for Patents
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INFORMATION DISCLOSURE STATEMENT

The information disclosure statement submitted herewith is being filed:

- ☒ Within three months of the filing date of the application or date of entry into the national stage of an international application or before the mailing date of a first Office action on the merits, whichever event occurs last. 37 C.F.R. section 1.97(b).

OR

- ☐ After three months of the filing date of this national application or the date of entry of the national stage as set forth in Section 1.491 in an international application or after the mailing date of the first Office action on the merits, whichever event occurred last but *before* the mailing date of either:

CERTIFICATION UNDER 37 C.F.R. SECTIONS 1.8(a) and 1.10*

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Carolyn Willey

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Practitioner's Docket No. MPI95-015P1RCPA1DV1M

- (1) a final action under Section 1.113,
- (2) a notice of allowance under Section 1.311, or
- (3) an action that otherwise closes prosecution in the application

whichever occurs first.

☐ Accompanying this transmittal is the fee set forth in 37 C.F.R. Section 1.17(p) for submission of an information disclosure statement under Section 1.97(c). (\$180.00).

☐ Each item of information contained in the information disclosure statement was cited in a communication from a foreign patent office in a counterpart foreign application not more than three months prior to the filing of the information disclosure statement. 37 C.F.R. Section 1.97(e)(1).

☐ No item of information contained in the information disclosure statement was cited in a communication from a foreign patent office in a counterpart foreign application and to the knowledge of the person signing the statement after making reasonable inquiry, was known to any individual designated in Section 1.56(c) more than three months prior to the filing of the information disclosure statement. 37 C.F.R. Section 1.97(e)(2).

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☐ The information disclosure statement transmitted herewith is being filed *after* a final action under Section 1.113, or a notice of allowance under Section 1.311, whichever occurs first, but before, or simultaneously with, the payment of the issue fee.

☐ In accordance with the requirements of 37 C.F.R. Section 1.97(d):

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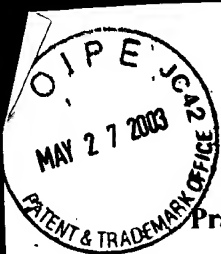
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<u>May 22, 2003</u>	MILLENNIUM PHARMACEUTICALS, INC. By <u>Tracy M. Sioussat</u> Tracy M. Sioussat Registration No. 50,609 75 Sidney Street Cambridge, MA 02139 Telephone - 617-374-7679 Facsimile - 617-551-8820
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Application No.: 09/801,089
Filed: March 8, 2001
For: MODULATION OF INTEGRIN-MEDIATED SIGNAL TRANSDUCTION

Group No.: 1644

Examiner: Ewoldt, Gerald R.

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Alexandria, VA 22313-1450

INFORMATION DISCLOSURE STATEMENT

List of Sections Forming Part of This Information Disclosure Statement

The following sections are being submitted for this Information Disclosure Statement:

1. [x] Preliminary Statements
2. [x] Forms PTO/SB/08A and 08B (substitute for Form PTO-1449)
3. [x] Identification of Prior Application in Which Listed Information Was Already Cited and for Which No Copies Are Submitted or Need Be Submitted
4. [x] Identification of Person(s) Making This Information Disclosure Statement

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Section 1. Preliminary statements

Applicants submit herewith patents, publications or other information, of which they are aware that they believe may be material to the examination of this application, and in respect of which, there may be a duty to disclose.

The filing of this information disclosure statement shall not be construed as a representation that a search has been made (37 C.F.R. section 1.97(g)), an admission that the information cited is, or is considered to be, material to patentability, or that no other material information exists.

The filing of this information disclosure statement shall not be construed as an admission against interest in any manner. Notice of January 9, 1992, 1135 O.G. 13-25, at 25.

Section 4. Identification of Prior Application in Which Listed Information Was Already Cited and for Which No Copies Are Submitted or Need Be Submitted

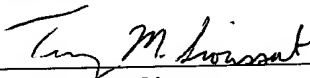
This application relies, under 35 U.S.C. section 120, on the earlier filing date of prior application SN.: 08/734,607 filed on October 18, 1995.

The references were submitted to, and/or cited by, the Office in the prior application(s) and, therefore, are not required to be provided in this application.

May 22, 2003

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By

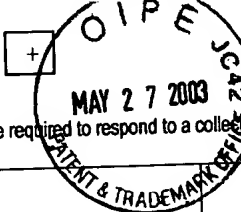


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INFORMATION DISCLOSURE STATEMENT BY APPLICANT

(use as many sheets as necessary)

Sheet 1 of 6

Complete if Known

Application Number	09/801,089
Filing Date	March 8, 2001
First Named Inventor	Phillips, David R.
Group Art Unit	1644
Examiner Name	Ewoldt, Gerald R.
Attorney Docket Number	MPI95-015P1

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		Mikayama, T., et al. Molecular Cloning and Functional Expression of CDNA Encoding Glycosylation Inhibiting Factor Proceedings of the National Academy of Sciences USA (90) 10056-10060 November 1993.	
		Scarborough, et al., Design of potent and specific integrin antagonists. Peptide antagonists with high specificity for glycoprotein IIb-IIIa, pp60c-src, pp62c-yes, and the p21ras GTPase-activating protein with the membrane skeleton. Journal of Biological Chemistry 268:1066-1073 (1993).	
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		Dorahy, et al., v Capture by chemical crosslinkers provides evidence that integrin alpha IIb beta 3 forms a complex with protein tyrosine kinases in intact platelets. Biochemistry Journal 309: 481-490 (1995).	
		Argaves, W.S., et al., Fibulin, a novel protein that interacts with the fibronectin receptor beta subunit cytoplasmic domain. Cell 58:623-629 (1989).	
		Bartfield, N.S., et al., The alpha v beta 3 integrin associates with a 190-kDa protein that is phosphorylated on tyrosine in response to platelet-derived growth factor. Journal of Biological Chemistry 268:17270-17276 (1993).	
		Chen, Y.P., et al., Ser-752-->Pro mutation in the cytoplasmic domain of integrin beta 3 subunit and defective activation of platelet integrin alpha IIb beta 3 (glycoprotein IIb-IIIa) in a variant of Glanzmann thrombasthenia. Proceedings of the National Academy of Sciences USA 89:10169-10173 (1992).	
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		Elmore, M. A., et al., Tyrosine-specific phosphorylation of gpIIa in platelet membranes. FEBS Letters 269:283-287 (1990).	
		Filardo, E.J., et al., Requirement of the NPXY motif in the integrin beta 3 subunit cytoplasmic tail for melanoma cell migration in vitro and in vivo. Journal of Cell Biology 130:441-450(1995).	

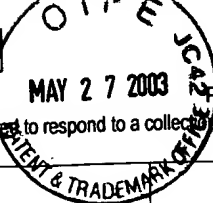
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INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)		Application Number	09/801,089
		Filing Date	March 8, 2001
		First Named Inventor	Phillips, David R.
		Group Art Unit	1644
		Examiner Name	Ewoldt, Gerald
		Attorney Docket Number	MPI95-015P1RCPA1DV1M
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		Findik, D., et al., Platelet membrane glycoproteins IIb and III a are substrates of purified pp60c-src protein tyrosine kinase. FEBS Letters 262:1-4 (1990).	
		Fitzgerald, L., et al., Protein sequence of endothelial glycoprotein IIIa derived from a cDNA clone. Identity with platelet glycoprotein IIIa and similarity to "integrin". Journal of Biological Chemistry 262:3936-3939 (1987).	
		Ginsberg, M.H., et al., Inside-out integrin signaling. Current Opinion in Cell Biology 4:766-771 (1992).	
		Ginsberg, M.H., et al., Platelet integrins. Thrombosis and Haemostasis 70:87-93 (1993).	
		Ginsberg, M.H., et al., Dynamic regulation of integrins. Stem Cells, 13:38-46 (1995).	
		Grinblat, Y., et al., Functions of the cytoplasmic domain of the beta PS integrin subunit during Drosophila development. Development 120:91-102 (1994).	
		Hannigan, G.E., et al., Regulation of cell adhesion and anchorage-dependent growth by a new beta 1-integrin-linked protein kinase. Nature 379:91-96 (1996).	
		Hayashi, Y., et al. Expression and function of chicken integrin beta 1 subunit and its cytoplasmic domain mutants in mouse NIH 3T3 cells. Journal of Cell Biology 110:175-184 (190).	
		Hibbs, M.L., et al., The cytoplasmic domain of the integrin lymphocyte function-associated antigen 1 beta subunit: sites required for binding to intercellular adhesion molecule 1 and the phorbol ester-stimulated phosphorylation site. Journal of Experimental Medicine 174:1227-1238 (1991).	

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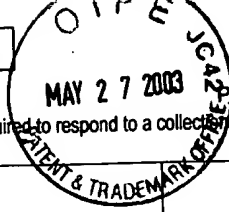
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Sheet 3 of 6

Application Number	09/801,089
Filing Date	March 8, 2001
First Named Inventor	Phillips, David R.
Group Art Unit	1644
Examiner Name	Ewoldt, Gerald
Attorney Docket Number	MPI95-015P1RCPA1DV1M

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		Hillary, C.A., et al., Phosphorylation of human platelet glycoprotein IIIa (GPIIIa). Disociation from fibrinogen receptor activation and phosphorylation of GPIIIa in vitro. Journal of Biological Chemistry 266:14663-14669 (1991).	
		Hirst, R., et al., Phosphorylation of the fibronectin receptor complex in cells transformed by oncogenes that encode tyrosine kinases. Proceedings of the National Academy of Sciences USA 83:6470-6474 (1986).	
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		Huang, M-M., et al., Adhesive ligand binding to integrin alpha IIb beta 3 stimulates tyrosine phosphorylation of novel protein substrates before phosphorylation of pp125FAK. Journal of Cell Biology 122:473-483 (1993).	
		Hughes, P.E., et al., The conserved membrane-proximal region of an integrin cytoplasmic domain specifies ligand binding affinity. Journal of Biological Chemistry 270:12411-12417 (1995).	
		Hynes, R.O., Integrins: versatility, modulation, and signaling in cell adhesion. Cell 69:11-25 (1992).	
		Johansson, M.W., et al., Altered localization and cytoplasmic domain-binding properties of tyrosine-phosphorylated beta 1 integrin. Journal of Cell Biology 126:1299-1309 (1994).	
		Juliano, R.L., et al., Signal transduction from the extracellular matrix. Cell Biology 120:577-585 (1993).	
		Kieffer, N., et al., Adhesive properties of the beta 3 integrins: comparison of GP IIb-IIIa and the vitronectin receptor individually expressed in human melanoma cells. Journal of Cell Biology 113:451-461 (1991).	

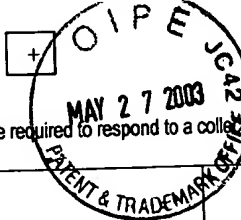
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Sheet 4 of 6

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Application Number 09/801,089
Filing Date March 8, 2001
First Named Inventor Phillips, David R.
Group Art Unit 1644
Examiner Name Ewoldt, JEFFREY
Attorney Docket Number MPI95-015P1RCPATB44A 1600/2900

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		LaFlamme, S.E., et al., Single subunit chimeric integrins as mimics and inhibitors of endogenous integrin functions in receptor localization, cell spreading and migration, and matrix assembly. Journal of Cell Biology 126:1287-1298 (1994).	
		Lanza, F., et al., cDNA cloning and expression of platelet p24/CD9. Evidence for a new family of multiple membrane-spanning proteins. Journal of Biological Chemistry. 266:10638-10645 (1991)	
		Luscinskas, et al. Integrins as dynamic regulators of vascular function. The FASEB Journal 8:929-938 (1994).	
		O'Toole, T.E., et al., Affinity modulation of the alpha IIb beta 3 integrin (platelet GPIIb-IIIa) is an intrinsic property of the receptor. Cell Regulation 1:883-893 (1990).	
		O'Toole, T.E., Regulation of integrin affinity states through an NPXY motif in the beta subunit cytoplasmic domain. Journal of Biological Chemistry 270:8553-8558 (1995).	
		Otey, C.A., et al., Mapping of the alpha-actinin binding site within the beta 1 integrin cytoplasmic domain. Journal of Biological Chemistry 268:21193-21197 (1993).	
		Parise, L.V., et al., Glycoprotein IIIa is phosphorylated in intact human platelets. Blood 75:2363-2368 (1990).	
		Phillips, D.R., et al. The platelet membrane glycoprotein IIb-IIIa complex. Blood 71:831-843 (1988).	
		Phillips, D.R., et al., GPIIb-IIIa: the responsive integrin. Cell 65:359-362 (1991).	

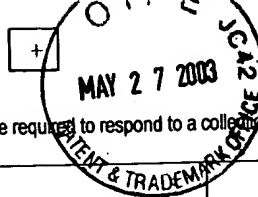
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				Examiner Name	Ewoldt, Gerald R.
				Attorney Docket Number	MPI95-015P1RCPA1DV1M

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		Reszka, A.A., Identification of amino acid sequences in the integrin beta 1 cytoplasmic domain implicated in cytoskeletal association. Journal of Cell Biology 117:1321-1330 (1992).	
		Rouslahti, E., Integrins. The Journal of Clinical Investigations 87:1-5 (1991).	
		Schaller, M.D., et al., Focal adhesion kinase and paxillin bind to peptides mimicking beta integrin cytoplasmic domains. Journal of Cell Biology 130:1181-1187 (1995).	
		Shattil, S.J., et al., Thrombosis and Haemostasis 73:1190 (1995).	
		Shattil, S.J., et al., Changes in the platelet membrane glycoprotein IIb/IIIa complex during platelet activation. Journal Biological Chemistry 260:11107-11114 (1985).	
		Smyth, S.S., et al., Regulation of vascular integrins. Blood 81:2827-2843 (1993).	
		Diamond, M.S., The dynamic regulation of integrin adhesiveness. Current Biology 4:506-517 (1994).	
		Tamkun, J.W., et al., Structure of integrin, a glycoprotein involved in the transmembrane linkage between fibronectin and actin. Cell 46:271-282 (1986).	
		Tapley, P., et al., Integrins isolated from Rous sarcoma virus-transformed chicken embryo fibroblasts. Oncogene 4:325-333 (1989).	

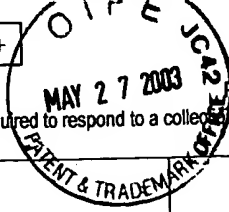
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		Filing Date	March 8, 2001
		First Named Inventor	Phillips, David R.
		Group Art Unit	1644
		Examiner Name	Ewoldt, Gerald R.
		Attorney Docket Number	MPI95-015P1RCPA1DV1M
Sheet	6	of	6

OTHER PRIOR ART -- NON PATENT LITERATURE DOCUMENTS			
Examiner Initials*	Cite No. ¹	Include the name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and-or country where published.	T ²
		Tcheng, J.E., et al., Multicenter, randomized, double-blind, placebo-controlled trial of the platelet integrin glycoprotein IIb/IIIa blocker Integrelin in elective coronary intervention. IMPACT Investigators. Circulation 91:2151-5157 (1995).	
		The EPIC Investigation, New England Journal of Medicine 330:956-961 (1994).	
		Vuori, K., et al., Mutation of the cytoplasmic domain of the integrin beta 3 subunit. Differential effects on cell spreading, recruitment to adhesion plaques, endocytosis, and phagocytosis. Journal of Biological Chemistry 270:9550-9557 (1995).	
		Ylanne, J., et al., Distinct functions of integrin alpha and beta subunit cytoplasmic domains in cell spreading and formation of focal adhesion. Journal of Cell Biology 122:223-233 (1993).	
		Lanza, Characterization of the human platelet glycoprotein IIIa gene. Comparison with the fibronectin receptor beta-subunit gene. Journal of Biological Chemistry, 265(30):18098-18103 (1990).	
		Lukashev, Disruption of integrin function and induction of tyrosine phosphorylation by the autonomously expressed beta 1 integrin cytoplasmic domain. Journal of Biological Chemistry, 269(28):18311-18314 (1994).	

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